

protein and which generates a signal in a manner dependent on the binding of the first binding partner to the protein, and a labelled second binding partner which binds to said protein in a manner independent of the conformational state of said protein and which generates a signal in a manner dependent on the binding of the first binding partner to the protein, wherein said protein and said labelled first and second binding partner are not covalently bound; and

- b) detecting said protein by the binding of at least one of said labelled first binding partner or said labelled second binding partner to said protein wherein detection of a signal generated by said labelled first binding partner and/or said labelled second binding partner is an indicator of the conformational state of said protein.

2. (Three Times Amended) A method for measuring the post-translational modifying activity of an enzyme, wherein the conformation of a protein is dependent upon the post-translational modification activity of the enzyme, the method comprising the steps of:

- a) contacting a protein comprising a site for post-translational modification with the enzyme;
- b) providing a labelled first binding partner which binds non-covalently to the protein in a manner dependent on the post-translational modification of the protein by the enzyme and which generates a signal in a manner dependent on said post-translational modification, and a second labelled binding partner which binds non-covalently to said protein and which generates a signal in a manner dependent on said post-translational modification;
- c) contacting the protein with the labelled first binding partner and the labelled second binding partner and detecting said protein by the binding

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to said protein of at least one of said labelled first binding partner and said labelled second binding partner, wherein detection of a signal generated by said labelled second binding partner and/or said labelled second binding partner indicates the post-translational modifying activity of the enzyme.

13. (Three Times Amended) The method of claim 1 or 2, further comprising the additional step of, after step (a), removing unbound labelled first binding partner to allow detection of the binding of the labelled first binding partner to the protein.

19. (Twice Amended) A kit for the determination of the conformational state of a protein in a sample, comprising:

- a) a labelled first binding partner which binds to the protein in a manner dependent on the conformational state of the protein and is detectable in a manner dependent on its binding to the protein and a labelled second binding partner which binds to the protein in a manner independent of the conformational state of said protein and which generates a signal in a manner dependent on the binding of the first binding partner to the protein, wherein said protein and said first binding partner and said second binding partner are not covalently bound; and
- b) packaging components.

20. (Twice Amended) A kit for the determination of the presence of a ligand for a protein in a sample, comprising:

- a) a protein which binds to the ligand the presence of which is to be determined and which undergoes a conformational change as a result of such binding;